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D (b) determining ^{the} a level of 6-thioguanine in said subject having said immune-mediated gastrointestinal disorder,

D wherein ^{the} a level of 6-thioguanine less than ~~a level~~ ^e ~~corresponding to~~ about 230 pmol per 8×10^8 red blood cells indicates a need to increase the amount of said drug subsequently administered to said subject and

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D wherein ^{the} a level of 6-thioguanine greater than ~~a level~~ ^e ~~corresponding to~~ about 400 pmol per 8×10^8 red blood cells indicates a need to decrease the amount of said drug subsequently administered to said subject.

7. A method of reducing toxicity associated with treatment of an immune-mediated gastrointestinal disorder, comprising:

(a) administering a drug providing 6-thioguanine to a subject having said immune-mediated gastrointestinal disorder;

CJD (b) determining ^{the} a level of 6-thioguanine in said subject having said immune-mediated gastrointestinal disorder; and

0 (c) determining ^{the} a level of 6-methyl-mercaptopurine in said subject having said immune-mediated gastrointestinal disorder,

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wherein ^{the} a level of 6-thioguanine greater than about 400 pmol per 8×10^8 red blood cells or ^{the} a level of 6-methyl-mercaptopurine greater than about 7000 pmol per 8×10^8 red blood cells indicates a need to decrease the amount of said drug subsequently administered to said subject, ~~thereby reducing toxicity associated with said drug treatment of said immune mediated gastrointestinal disorder.~~

15. A method of optimizing therapeutic efficacy and reducing toxicity associated with treatment of an immune-mediated gastrointestinal disorder, comprising:

(a) administering a drug providing 6-thioguanine to a subject having said immune-mediated gastrointestinal disorder;

(b) determining ^{the} a level of 6-thioguanine in said subject having said immune-mediated gastrointestinal disorder; and

(c) determining ^{the} a level of 6-methyl-mercaptopurine in said subject having said immune-mediated gastrointestinal disorder,

wherein ^{the} a level of 6-thioguanine less than about 230 pmol per 8×10^8 red blood cells indicates a need to increase the amount of said drug subsequently administered to said subject, ~~thereby increasing therapeutic efficacy.~~

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D wherein ^{the} a level of 6-thioguanine greater than about 400 pmol per 8×10^8 red blood cells indicates a need to decrease the amount of said drug subsequently administered to said subject ✓

0 ~~thereby reducing toxicity associated with said drug treatment of~~
~~said immune-mediated gastrointestinal disorder, and~~

C30 wherein ^{the} a level of 6-methyl-mercaptopurine greater than about 7000 pmol per 8×10^8 red blood cells indicates a need to decrease the amount of said drug subsequently administered to said subject ✓

0 ~~thereby reducing toxicity associated with said drug~~
~~treatment of said immune-mediated gastrointestinal disorder.~~

22 30. A method of optimizing therapeutic efficacy of treatment of a non-IBD autoimmune disease, comprising:

(a) administering a drug providing 6-thioguanine to a subject having said non-IBD autoimmune disease; and

C4 0 (b) determining ^{the} level of 6-thioguanine in said subject having said non-IBD autoimmune disease,

0 wherein ^{the} a level of 6-thioguanine less than about 230 pmol per 8×10^8 red blood cells indicates a need to increase the amount of 6-mercaptopurine drug subsequently administered to said subject and

D wherein ^{the} a level of 6-thioguanine greater than about 400 pmol per 8×10^8 red blood cells indicates a need to decrease the

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C4 amount of 6-mercaptopurine drug subsequently administered to said subject.

25. A method of optimizing therapeutic efficacy and reducing toxicity associated with treatment of an immune-mediated gastrointestinal disorder, comprising:

(a) administering a drug providing 6-thioguanine to a subject having said immune-mediated gastrointestinal disorder;

D (b) determining ^{the} ~~a~~ level of 6-thioguanine in said subject having said immune-mediated gastrointestinal disorder; and

C4 (c) determining ^{the} ~~a~~ level of 6-methyl-mercaptopurine in said subject having said immune-mediated gastrointestinal disorder,

D wherein ^{the} ~~a~~ level of 6-thioguanine less than about 230 pmol per 8×10^8 red blood cells indicates a need to increase the amount of said drug subsequently administered to said subject, ~~thereby increasing therapeutic efficacy,~~ and

D wherein ^{the} ~~a~~ level of 6-thioguanine greater than about 400 pmol per 8×10^8 red blood cells or a level of 6-methyl-mercaptopurine greater than about 7000 pmol per 8×10^8 red blood cells indicates a need to decrease the amount of said drug subsequently administered to said subject, ~~thereby reducing~~

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CS cont ~~toxicity associated with said drug treatment of said
immune-mediated gastrointestinal disorder.~~

52. A method of optimizing therapeutic efficacy of treatment of a non-IBD autoimmune disease, comprising:

(a) administering a drug providing 6-thioguanine to a subject having said non-IBD autoimmune disease;

(b) determining a level of 6-thioguanine in said subject having said non-IBD autoimmune disease; and

Cu (c) determining a level of 6-methyl-mercaptopurine in said subject having said non-IBD autoimmune disease,

wherein a level of 6-thioguanine less than about 230 pmol per 8×10^8 red blood cells indicates a need to increase the amount of said drug subsequently administered to said subject and

wherein a level of 6-thioguanine greater than about 400 pmol per 8×10^8 red blood cells or a level of 6-methyl-mercaptopurine greater than about 7000 pmol per 8×10^8 red blood cells indicates a need to decrease the amount of said drug subsequently administered to said subject, thereby reducing toxicity associated with said drug treatment of said non-IBD autoimmune disease.

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Please add the following new claims:

37/55. (New) A method of optimizing therapeutic efficacy and reducing toxicity associated with treatment of an immune-mediated gastrointestinal disorder, comprising:

(a) administering a drug selected from the group consisting of 6-mercaptopurine, azathioprine, 6-thioguanine, and 6-methylmercaptopurine to a subject having said immune-mediated gastrointestinal disorder; and

(b) determining ^{the} level of 6-thioguanine or 6-methyl-mercaptopurine in said subject having said immune-mediated gastrointestinal disorder;

wherein ^{the} level of 6-thioguanine less than about 230 pmol per 8×10^8 red blood cells indicates a need to increase the amount of said drug subsequently administered to said subject, ~~thereby increasing therapeutic efficacy,~~ and

wherein ^{the} level of 6-thioguanine greater than about 400 pmol per 8×10^8 red blood cells or a level of 6-methyl-mercaptopurine greater than about 7000 pmol per 8×10^8 red blood cells indicates a need to decrease the amount of said drug subsequently administered to said subject, ~~thereby reducing toxicity associated with said drug treatment of said immune mediated gastrointestinal disorder.~~

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~~38~~ 56. (New) The method of claim ~~37~~ 55, wherein said drug is 6-mercaptopurine.

~~39~~ 57. (New) The method of claim ~~37~~ 55, wherein said drug is azathioprine.

~~40~~ 58. (New) The method of claim ~~37~~ 55, wherein said immune-mediated gastrointestinal disorder is inflammatory bowel disease (IBD).

~~41~~ 59. (New) The method of claim ~~40~~ 58, wherein said subject having IBD is a pediatric subject.

~~42~~ 60. (New) The method of claim ~~37~~ 55, wherein said immune-mediated gastrointestinal disorder is selected from the group consisting of lymphocytic colitis, microscopic colitis, collagenous colitis, autoimmune enteropathy, allergic gastrointestinal disease and eosinophilic gastrointestinal disease.

~~43~~ 61. (New) The method of claim ~~37~~ 55, wherein said level of 6-thioguanine and said level of 6-methyl-mercaptopurine each is determined in red blood cells.

~~44~~ 62. (New) The method of claim ~~43~~ 61, wherein said level is determined using high pressure liquid chromatography.

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63. (New) The method of claim 37, wherein said toxicity associated with said drug treatment is selected from the group consisting of hepatic toxicity and hematologic toxicity.

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64. (New) A method of optimizing therapeutic efficacy and reducing toxicity associated with treatment of an immune-mediated gastrointestinal disorder, comprising:

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D (a) determining ^{the} a level of 6-thioguanine or 6-methyl-mercaptopurine in a subject administered a drug selected from the group consisting of 6-mercaptopurine, azathioprine, 6-thioguanine, and 6-methylmercaptoriboside, said subject having said immune-mediated gastrointestinal disorder;

D wherein ^{the} a level of 6-thioguanine less than about 230 pmol per 8×10^8 red blood cells indicates a need to increase the amount of said drug subsequently administered to said subject, ~~thereby increasing therapeutic efficacy,~~ and

D wherein ^{the} a level of 6-thioguanine greater than about 400 pmol per 8×10^8 red blood cells or a level of 6-methyl-mercaptopurine greater than about 7000 pmol per 8×10^8 red blood cells indicates a need to decrease the amount of said drug subsequently administered to said subject, ~~thereby reducing toxicity associated with said drug treatment of said immune-mediated gastrointestinal disorder.~~

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65. (New) The method of claim 46, wherein said drug is 6-mercaptopurine.

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~~48~~
~~66~~. (New) The method of claim ~~64~~, wherein said drug is azathioprine.

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~~67~~. (New) The method of claim ~~64~~, wherein said immune-mediated gastrointestinal disorder is IBD.

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~~68~~. (New) The method of claim ~~65~~, wherein said subject having IBD is a pediatric subject.

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~~69~~. (New) The method of claim ~~64~~, wherein said immune-mediated gastrointestinal disorder is selected from the group consisting of lymphocytic colitis, microscopic colitis, collagenous colitis, autoimmune enteropathy, allergic gastrointestinal disease and eosinophilic gastrointestinal disease.

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~~70~~. (New) The method of claim ~~64~~, wherein said level of 6-thioguanine and said level of 6-methyl-mercaptopurine each is determined in red blood cells.

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~~71~~. (New) The method of claim ~~70~~, wherein said level is determined using high pressure liquid chromatography.

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~~72~~. (New) The method of claim ~~64~~, wherein said toxicity associated with said drug treatment is selected from the group consisting of hepatic toxicity and hematologic toxicity.

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